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FILE 'USPATFULL' ENTERED AT 11:23:11 ON 17 AUG 2004
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FILE 'BIOSIS' ENTERED AT 11:23:11 ON 17 AUG 2004
COPYRIGHT (C) 2004 BIOLOGICAL ABSTRACTS INC.(R)
=> s rna and (2()o()methyl)
L1
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=> s 12 and py<1997
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     ANSWER 1 OF 27 USPATFULL ON STN
                          2001:220860 USPATFULL
ACCESSION NUMBER:
TITLE:
                          Enzymatic DNA molecules
                         Joyce, Gerald F., Encinitas, CA, United States
Breaker, Ronald R., Guilford, CT, United States
INVENTOR(S):
PATENT ASSIGNEE(S):
                         The Scripps Research Institute, La Jolla, CA, United
                          States (U.S. corporation)
                              NUMBER
                                            KIND
                                                     DATE
PATENT INFORMATION:
                         US 6326174
                                                   20011204
                         wo 9617086
                                                   19960606
                                                                         <--
APPLICATION INFO.:
                         US 1997-849567
                                                   19970825
                                                              (8)
                                                   19951201
                         wo 1995-us15580
                                                   19970825
                                                              PCT 371 date
                                                   19970825 PCT 102(e) date
DOCUMENT TYPE:
                         Utility
FILE SEGMENT:
                         GRANTED
PRIMARY EXAMINER:
                         Schwartzman, Robert A.
LEGAL REPRESENTATIVE:
                         Fitting, Thomas, Holmes, Emily
NUMBER OF CLAIMS:
                         68
EXEMPLARY CLAIM:
NUMBER OF DRAWINGS:
                         11 Drawing Figure(s); 9 Drawing Page(s)
LINE COUNT:
                         2703
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention discloses deoxyribonucleic acid enzymes--catalytic
       or enzymatic DNA molecules--capable of cleaving nucleic acid sequences
       or_molecules, particularly ***RNA*** , in a site-specific manner, as
       well as compositions including same. Methods of making and using the disclosed enzymes and compositions are also disclosed.
```

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2001:147670 USPATFULL ACCESSION NUMBER:

Compositions and methods for treatment of hepatitis C TITLE:

virus-associated diseases

INVENTOR(S):

Anderson, Kevin P., Carlsbad, CA, United States Hanecak, Ronnie C., San Clemente, CA, United States

Hoshiko, Kazuya, Koshi-machi, Japan Nozaki, Chikateru, Kumamoto, Japan Nishihara, Tsukasa, Kumamoto, Japan Nakatake, Hiroshi, Kikuyo-machi, Japan Hamada, Fukusaburo, Nishigoshi-machi, Japan

Eto, Tatsuo, Ohzu-machi, Japan

Furukawa, Shinichi, Koshi-machi, Japan

PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., Carlsbad, CA, United States

(U.S. corporation)

NUMBER KIND DATE ______ us 6284458 PATENT INFORMATION: 20010904 в1 wo 9405813 19940317 <-us 1995-397220 APPLICATION INFO.: 19950309 (8) WO 1993-JP1293 19930910 19950309 PCT 371 date 19950309 PCT 102(e) date

NUMBER DATE

PRIORITY INFORMATION: JP 1993-87195 19930414

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED PRIMARY EXAMINER: Wang, Andrew

LEGAL REPRESENTATIVE: Licata & Tyrrell P.C.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 9 Drawing Figure(s); 9 Drawing Page(s)

LINE COUNT: 1208

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Antisense oligonucleotides are provided which are complementary to and AB hybridizable with at least a portion of HCV ***RNA*** and which are capable of inhibiting the function of the HCV ***RNA***. These oligonucleotides can be administered to inhibit the activity of Hepatitis C virus in vivo or in vitro. These compounds can be used either prophylactically or therapeutically to reduce the severity of diseases associated with Hepatitis C virus, and for diagnosis and detection of HCV and HCV-associated diseases. Methods of using these compounds are also disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 3 OF 27 USPATFULL on STN L4

2001:142075 USPATFULL ACCESSION NUMBER:

TITLE: High affinity nucleic acid ligands to lectins INVENTOR(S): Parma, David H, Boulder, CO, United States Hicke, Brian, Boulder, CO, United States Bridonneau, Philippe, Boulder, CO, United States Gold, Larry, Boulder, CO, United States

PATENT ASSIGNEE(S): Gilead Sciences, Inc., Foster City, CA, United States

(U.S. corporation)

NUMBER KIND DATE US 6280932 B1 PATENT INFORMATION: 20010828 wo 9640703 19961219 <--US 1997-952793 APPLICATION INFO.: 19971121 (8) wo 1996-us9455 19960605 19971121 PCT 371 date 19971121 PCT 102(e) date

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1995-479724,

on 7 Jun 1995, now patented, Pat. No. US 5780228 Continuation-in-part of Ser. No. US 1995-472256, filed on 7 Jun 1995, now patented, Pat. No. US 6001988 Continuation-in-part of Ser. No. US 1995-472255, filed on 7 Jun 1995, now patented, Pat. No. US 5766853 Continuation-in-part of Ser. No. US 1995-477829, filed on 7 Jun 1995, now abandoned Continuation-in-part of Ser. No. US 1991-714131 filed on 10 Jun 1991

Ser. No. US 1991-714131, filed on 10 Jun 1991, now patented, Pat. No. US 5475096 Continuation-in-part of abandoned

DOCUMENT TYPE: FILE SEGMENT:

Utility GRANTED Riley, Jezia

PRIMARY EXAMINER: LEGAL REPRESENTATIVE:

Swanson & Bratschun, L.L.C.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

50

NUMBER OF DRAWINGS:

24 Drawing Figure(s); 19 Drawing Page(s)

LINE COUNT:

4625

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention discloses high-affinity oligonucleotide ligands to AB lectins, specifically nucleic acid ligands having the ability to bind to the lectins, wheat germ agglutinin, L-selectin and P-selectin. Also

disclosed are the methods for obtaining such ligands.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

ANSWER 4 OF 27 USPATFULL on STN

TITLE:

2001:136439 USPATFULL

INVENTOR(S):

Optimized minizymes and miniribozymes and uses thereof

McCall, Maxine J., Putney, Australia Hendry, Philip, Leichhardt, Australia

PATENT ASSIGNEE(S):

Lockett, Trevor, Denistone, Australia Commonwealth Scientific and Industrial Research

Organization, Parkville, Australia (non-U.S.

corporation)

NUMBER KIND DATE

PATENT INFORMATION:

US 6277634 20010821 wo 9640906 19961219

APPLICATION INFO.:

us 1998-973568 19980518 (8)

WO 1996-AU343 19960607

<--

RELATED APPLN. INFO.:

19980515 PCT 371 date 19980515 PCT 102(e) date Continuation of Ser. No. US 1995-574396, filed on 18

Dec 1995, now patented, Pat. No. US 6001648 Continuation-in-part of Ser. No. US 1995-488181,

on 7 Jun 1995, now patented, Pat. No. US 6004806

DOCUMENT TYPE: FILE SEGMENT:

Utility GRANTED

2590

PRIMARY EXAMINER:

McGarry, Sean

LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS:

White, John P. Cooper & Dunham LLP

EXEMPLARY CLAIM:

46

NUMBER OF DRAWINGS:

5 Drawing Figure(s); 4 Drawing Page(s)

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention is directed to improved catalytic compounds, minizymes and miniribozymes, capable of hybridizing with a target ***RNA*** be cleaved. The minizymes and miniribozymes and compositions of the present invention may be used in vitro or in vivo. They may be used as diagnostic or therapeutic agents.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

14 ANSWER 5 OF 27 USPATFULL on STN

ACCESSION NUMBER:

2000:153836 USPATFULL Nucleic acid ligand complexes

TITLE: INVENTOR(S):

Gold, Larry, Boulder, CO, United States Schmidt, Paul G, Niwot, CO, United States Janjic, Nebojsa, Boulder, CO, United States NeXstar Pharmaceuticals, Inc., Boulder, CO, United

PATENT ASSIGNEE(S):

States (U.S. corporation)

NUMBER KIND DATE US 6147204 PATENT INFORMATION: 20001114 wo 9634876 19961107 <--APPLICATION INFO.: 19971028 US 1997-945604 (8) wo 1996-us6171 19960502 19971028

PCT 371 date PCT 102(e) date 19971028

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1995-434465, filed on 4 May 1995, now patented, Pat. No. US 6011020 And a continuation-in-part of Ser. No. US 1995-464443, filed

continuation-in-part of Ser. No. US 1991-714131, filed on 10 Jun 1991, now patented, Pat. No. US 5475096 which is a continuation-in-part of Ser. No. US 1990-536428,

filed on 11 Jun 1990, now abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER:

Zitomer, Stephanie Swanson & Bratschun, L.L.C. LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: **EXEMPLARY CLAIM:**

NUMBER OF DRAWINGS: 36 Drawing Figure(s); 34 Drawing Page(s)

LINE COUNT: 2756

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention discloses a method for preparing a therapeutic or

diagnostic complex comprised of a nucleic acid ligand and a lipophilic compound or non-immunogenic, high molecular weight compound by identifying a nucleic acid ligand by SELEX methodology and associating the nucleic acid ligand the nucleic acid ligand population compound or a non-immunogenic, high molecular weight compound. The invention further discloses

complexes comprising one or more nucleic acid ligands in association with a lipophilic compound or non-immunogenic, high molecular weight

compound.

AB

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 6 OF 27 USPATFULL on STN

ACCESSION NUMBER: 2000:125209 USPATFULL

TITLE: Oligomeric compounds having nitrogen-containing

linkages

INVENTOR(S): Cook, Phillip Dan, Vista, CA, United States

Sanghvi, Yogesh S., San Marcos, CA, United States

Kung, Pei Pei, Carlsbad, CA, United States

PATENT ASSIGNEE(S): ISIS Pharmaceuticals, Inc., Carlsbad, CA, United States

(U.S. corporation)

	NUMBER	KIND DATE	
B4====================================			
PATENT INFORMATION:	US 6121433	20000919	
	wo 9518623	19950713	<
APPLICATION INFO.:	us 1996-669300	19960808	(8)
	wo 1995-us350	19950111	
		19960808	PCT 371 date

19960808 PCT 102(e) date Continuation-in-part of Ser. No. US 1994-180124, filed on 11 Jan 1994, now patented, Pat. No. US 5783682 And a continuation-in-part of Ser. No. US 1993-39979, filed RELATED APPLN. INFO.:

on 30 Mar 1993, now abandoned And a continuation-in-part of Ser. No. US 1993-39846, filed

on 30 Mar 1993, now abandoned And a

continuation-in-part of Ser. No. US 1993-40933. filed

on 31 Mar 1993, now abandoned And a continuation-in-part of Ser. No. US 1993-40903, filed on 31 Mar 1993, now patented, Pat. No. US 5386023 And a continuation-in-part of Ser. No. US 1993-40526, filed on 31 Mar 1993, now patented, Pat. No. US 5489677 which is a continuation-in-part of Ser. No. WO 1992-US4294, filed on 21 May 1992 And a continuation-in-part of Ser.

filed on 21 May 1992 And a continuation-in-part of Ser No. US 1992-903160, filed on 24 Jun 1992, now abandoned

which is a continuation-in-part of Ser. No. US

1991-703619, filed on 21 May 1991, now patented, Pat. No. US 5378825 which is a continuation-in-part of Ser. No. US 1990-566836, filed on 13 Aug 1990, now patented, Pat. No. US 5223618 And a continuation-in-part of Ser. No. US 1990-558663, filed on 27 Jul 1990, now patented, Pat. No. US 523663, filed on 27 Jul 1990, now patented,

Pat. No. US 5138045

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Marschel, Ardin H.

ASSISTANT EXAMINER: Riley, Jezia

LEGAL REPRESENTATIVE: Woodcock Washburn Kurtz Mackiewicz & Norris LLP

NUMBER OF CLAIMS: 13 **EXEMPLARY CLAIM:**

NUMBER OF DRAWINGS: 14 Drawing Figure(s); 14 Drawing Page(s)

LINE COUNT: 3461

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Novel compounds and libraries of compounds based on nitrogen atoms that

functional groups, that are attached to the nitrogen atoms, to the spanner groups or to both the nitrogen atoms and the spanner groups to render the compounds and libraries of such compounds with diverse properties.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 7 OF 27 USPATFULL ON STN

ACCESSION NUMBER: 2000:117501 USPATFULL

Systematic evolution of ligands by exponential TITLE:

enrichment: tissue selex

INVENTOR(S): Jensen, Kirk B, New York, NY, United States Chen, Hang, San Francisco, CA, United States

Morris, Kevin N., Goldegg, Austria
Stephens, Andrew, Boulder, CO, United States
Gold, Larry, Boulder, CO, United States
NeXstar Pharmaceuticals, Inc., Boulder, CO, United
States (U.S. corporation)

	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 6114120 WO 9634875		20000905 19961107	<
APPLICATION INFO.:	US 1997-945909 WO 1996-US6060		19971028 19960501	(8)
			19971028	PCT 371 date

19971028 PCT 102(e) date Continuation-in-part of Ser. No. US 1995-434425, filed on 3 May 1995, now patented, Pat. No. US 5789157 And a RELATED APPLN. INFO.: continuation-in-part of Ser. No. US 1995-437667, on 3 May 1995, now patented, Pat. No. US 5864026 And a

continuation-in-part of Ser. No. US 1995-434001, filed on 3 May 1995, now patented, Pat. No. US 5712375 And a continuation-in-part of Ser. No. US 1995-433585, filed on 3 May 1995, now patented, Pat. No. US 5763566

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Zitomer, Stephanie Swanson & Bratschun LLC LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 10

PATENT ASSIGNEE(S):

NUMBER OF DRAWINGS: 7 Drawing Figure(s); 7 Drawing Page(s)

LINE COUNT: 3551

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A new class of nucleic acid compounds, referred to as nucleic acid ligands, have been shown to exist that have a specific binding affinity AΒ for three dimensional molecular targets, including cell surface macromolecules. The nucleic acid ligands are identified by the method of the invention referred to as the Systematic Evolution of Ligands by EXponential enrichment (SELEX), wherein a candidate mixture of nucleic acids are iteratively enriched and the high affinity nucleic acids are amplified for further partitioning. The major acid ligands are useful in capturing target cells.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 8 OF 27 USPATFULL on STN

PATENT ASSIGNEE(S):

ACCESSION NUMBER: 2000:109986 USPATFULL

TITLE: Nitrogenous macrocyclic compounds

INVENTOR(S):

Cook, Phillip Dan, Escondido, CA, United States
An, Haoyun, Encinitas, CA, United States
Guinosso, Charles J., Vista, CA, United States
Kung, Pei-Pei, Leucadia, CA, United States
Fraser, Allister S., San Marcos, CA, United States
Isis Pharmaceuticals, Inc., Carlsbad, CA, United States

(U.S. corporation)

	NUMBER	KIND DATE	
PATENT INFORMATION:	US 6107482 WO 9630377	20000822 19961003	<
APPLICATION INFO.:	US 1997-913664 WO 1996-US4215	19970919 19960327	•
			PCT 371 date
RELATED APPLN. INFO.:	Continuation-in-	19970919 Dart of Ser. No.	PCT 102(e) date US 1995-461728, filed

No. US 1995-410703, filed on 27 Mar 1995

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Raymond, Richard L.

LEGAL REPRESENTATIVE: Woodcock Washburn Kurtz Mackiewicz & Norris LLP

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

AB

NUMBER OF DRAWINGS: 15 Drawing Figure(s); 15 Drawing Page(s)

LINE COUNT: 4599

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Novel macrocyclic compounds are constructed to include large cyclic structures that are interrupted by at least one ring system. Each interrupting ring system includes two bridgehead atoms. Bridgehead atoms are bonded to one or more bridges that interconnect one or more ring systems thereby forming a large cyclic structure. Located in each bridge are two or more nitrogenous moieties that are derivatized with chemical functional groups. The ring systems can include further nitrogenous moieties, either as ring atoms or on pendant groups attached to the ring. These can also be derivatized with chemical functional groups. The totality of the chemical functional groups imparts certain conformational and other properties to the macrocyclic compounds. In accordance with certain embodiments of the invention, libraries of such macrocyclic compounds are prepared utilizing permutations and combinations of the chemical functional groups and the nitrogenous moieties to build complexity into the library.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 9 OF 27 USPATFULL on STN

ACCESSION NUMBER: 2000:12593 USPATFULL

TITLE: Nucleic acid ligands that bind to and inhibit DNA

polymerases

INVENTOR(S): Gold, Larry, Boulder, CO, United States

Javasena, Sumedha, Boulder, CO, United States

NeXstar Pharmaceuticals, Inc., Boulder, CO, United PATENT ASSIGNEE(S):

States (U.S. corporation)

NUMBER KIND DATE US 6020130 PATENT INFORMATION: 20000201 wo 9641010 19961219 APPLICATION INFO.: US 1997-945734 19971028 (8) wo 1996-us9451 19960605 19971028 19971028 PCT 371 date PCT 102(e) date

Continuation-in-part of Ser. No. US 1995-487426, filed on 7 Jun 1995, now patented, Pat. No. US 5763173 Ser. No. Ser. No. US 1995-487720, filed on 7 Jun 1995, now

patented, Pat. No. US 5874557 And Ser. No. US 1995-484557, filed on 7 Jun 1995, now patented, Pat.

No. US 5693502 Utility

DOCUMENT TYPE: FILE SEGMENT: Granted

PRIMARY EXAMINER: Zitomer, Stephanie Swanson & Bratschun LLC LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: 17 **EXEMPLARY CLAIM:**

RELATED APPLN. INFO.:

NUMBER OF DRAWINGS: 35 Drawing Figure(s); 17 Drawing Page(s)

LINE COUNT: 2374

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention discloses high-affinity oligonucleotide ligands to the thermostable Taq polymerase and Tth polymerase. Specifically, this invention discloses DNA ligands having the ability to bind to the Taq and Tth polymerases and the methods for obtaining such ligands. The ligands are capable of inhibiting polymerases at ambient temperatures.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 10 OF 27 USPATFULL on STN

ACCESSION NUMBER: 1999:37288 USPATFULL

TITLE:

Phosphate linked oligomers
Cook, Phillip Dan, Vista, CA, United States
Acevedo, Oscar L., San Diego, CA, United States INVENTOR(S): Davis, Peter W., Carlsbad, CA, United States Ecker, David J., Encinitas, CA, United States

Hebert, Normand, Cardiff, CA, United States

(U.S. corporation) NUMBER KIND DATE PATENT INFORMATION: US 5886177 19990323 wo 9518820 19950713 <--US 1996-669506 APPLICATION INFO.: 19960808 (8) wo 1995-us449 19950111 19960808 PCT 371 date 19960808 PCT 102(e) date RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1994-179970, filed on 11 Jan 1994 DOCUMENT TYPE: Utility FILE SEGMENT: Granted PRIMARY EXAMINER: Marschel, Ardin н. Woodcock Washburn Kurtz Mackiewicz & Norris, LLP LEGAL REPRESENTATIVE: NUMBER OF CLAIMS: 22 EXEMPLARY CLAIM: NUMBER OF DRAWINGS: 4 Drawing Figure(s); 2 Drawing Page(s) LINE COUNT: 3915 CAS INDEXING IS AVAILABLE FOR THIS PATENT. Novel ethylene glycol compounds bearing various functional groups are used to prepare oligomeric structures. The ethylene glycol monomers can AB be joined via standard phosphate linkages including phosphorothioate, phosphodiester, and phosphoramidate linkages. Useful functional groups include nucleobases as well as polar groups, hydrophobic groups, ionic groups, aromatic groups and/or groups that participate in hydrogen-bonding. CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 11 OF 27 USPATFULL ON STN ACCESSION NUMBER: 1998:14925 USPATFULL TITLE: Phosphoramidate and phosphorothiomidate oligomeric compounds

INVENTOR(\$): Cook, Phillip Dan, Vista, CA, United States

Acevedo, Oscar, San Diego, CA, United States Hebert, Normand, Cardiff, CA, United States

PATENT ASSIGNEE(S): ISIS Pharmaceuticals, Inc., Carlsbad, CA, United States

(U.S. corporation)

NUMBER KIND DATE US 5717083 PATENT INFORMATION: 19980210 wo 9523160 19950831 APPLICATION INFO.: US 1996-693112 19960819 (8) wo 1995-us2267 19950223 19960819 PCT 371 date PCT 102(e) date 19960819 RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1994-200638,

on 23 Feb 1994, now patented, Pat. No. US 5637684

DOCUMENT TYPE: Utility FILE SEGMENT: Granted PRIMARY EXAMINER: Guzo, David

LEGAL REPRESENTATIVE: Woodcock Washburn Kurtz Mackiewicz & Norris LLP

NUMBER OF CLAIMS: 40 EXEMPLARY CLAIM: LINE COUNT: 2743

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Compounds are provided having structure (I), wherein the L groups are backbone segments, the Y and T groups are functional groups for interacting with target molecules of interest, the X groups are oxygen or sulfur and the E groups are H, conjugate groups or intermediate groups used during the synthesis of the compounds and wherein the AΒ compounds are prepared using H phosphonate type chemistry wherein the functional groups are added during an oxidation step or during a coupling step. ##STR1##

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 12 OF 27 USPATFULL on STN

ACCESSION NUMBER: 1998:12145 USPATFULL

TITLE: Pyrrolidine-containing monomers and oligomers INVENTOR(S): Acevedo, Oscar L., San Diego, CA, United States Hebert, Normand, Cardiff, CA, United States

PATENT ASSIGNEE(S): ISIS Pharmaceuticals, Inc., Carlsbad, CA, United States

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NUMBER
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 PATENT INFORMATION:
                           US 5714606
                                                      19980203
                                                      19950713
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                           US 1996-669505
APPLICATION INFO.:
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                           wo 1995-us356
                                                      19950111
                                                      19960815
                                                                PCT 371 date
PCT 102(e) date
                                                      19960815
                           Continuation-in-part of Ser. No. US 1994-180134, filed
RELATED APPLN. INFO.:
                           on 11 Jan 1994, now patented, Pat. No. US 5519134
DOCUMENT TYPE:
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FILE SEGMENT:
                           Granted
PRIMARY EXAMINER:
                           McKane, Joseph
LEGAL REPRESENTATIVE:
                           Woodcock Washburn Kurtz Mackiewicz & Norris, LLP
NUMBER OF CLAIMS:
                           26
EXEMPLARY CLAIM:
                           1
NUMBER OF DRAWINGS:
                           1 Drawing Figure(s); 1 Drawing Page(s)
LINE COUNT:
                           2992
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB
        The invention relates to pyrrolidine monomeric units and to oligomers
        which are joined via phosphate linkages, including phosphorothioate,
        phosphodiester and phosphoramidate linkages.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
      ANSWER 13 OF 27 CAPLUS COPYRIGHT 2004 ACS on STN
                            1997:231448 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                            126:288105
TITLE:
                            Ribozymes cleaving interleukin-5 mRNA for treatment
                            and diagnosis of asthma and other inflammatory
                            disorders
INVENTOR(S):
                            Sullivan, Sean; Draper, Kenneth G.; McSwiggen, James;
                            Stinchcomb, Dan T.
                            Ribozyme Pharmaceuticals, Inc., USA
U.S., 145 pp., Cont.-in-part of U.S. Ser. No. 989,849,
PATENT ASSIGNEE(S):
SOURCE:
                            abandoned.
                            CODEN: USXXAM
DOCUMENT TYPE:
                            Patent
LANGUAGE:
                            English
FAMILY ACC. NUM. COUNT:
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PATENT INFORMATION:
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170 A2 20021023 EP
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392

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490 A 19970401 US 1995-434503 19950504
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                                                 US 1992-987132
                                                                       Α
                                                                          19921207
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US 1992-989848

EP 1993-918144

19921207

A3 19930702

Α

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us 1994-218934
                         19940329
US 1994-222795
                         19940404
US 1994-224483
                         19940407
                     Α
us 1994-227958
                         19940415
US 1994-228041
                         19940415
US 1994-245736
                     Α
                         19940518
   1994-271280
                         19940706
US
                     Α
US
   1994-291932
                         19940815
                     Α
   1994-291433
US
                         19940816
US 1994-292620
                     Α
                         19940817
US 1994-293520
                     Α
                        19940819
us 1994-300000
                        19940902
                     Α
US 1994-303039
                     Α
                         19940908
us 1994-311486
                     Α
                         19940923
US
   1994-311749
                     Α
                        19940923
US
   1994-314397
                     Α
                        19940928
US
   1994-316771
                     Α
                        19941003
   1994-319492
US
                        19941007
                     Α
US
   1994-321993
                        19941011
US 1994-334847
                        19941104
us 1994-337608
                        19941110
US 1994-345516
                        19941128
US 1994-357577
                        19941216
US 1994-363233
                        19941223
US
   1995-380734
                        19950130
   1995-909920
EΡ
                     A3 19950223
WO 1995-IB156
                        19950223
                     W
AU 1995-26422
                     A3 19950518
US 1995-475460
                        19950607
                     Α
US 1995-483715
                        19950607
                     Α
US 1995-484607
                     Α
                        19950607
US 1996-623891
                     Α
                        19960325
AU 1996-61744
                     A3 19960603
AU 1996-76662
                     A3 19961025
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Ribozymes that cleave the mRNA of interleukin 5 are described for use in the therapeutic control of interleukin levels in the treatment of asthma AB and other inflammatory_diseases. Interleukin 5 levels are shown to be raised in bronchoalveolar lavage and lung biopsies of asthma patients, implying a role for helper T-cells in the inflammatory response.

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ANSWER 14 OF 27
                 USPATFULL on STN
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ACCESSION NUMBER:

TITLE:

96:111555 USPATFULL

INVENTOR(S):

Optimized catalytic DNA-cleaving ribozymes

PATENT ASSIGNEE(S):

Joyce, Gerald F., Encinitas, CA, United States The Scripps Research Institute, La Jolla, CA, United

States (U.S. corporation)

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NUMBER
                                             KIND
                                                      DATE
                          US 5580967
US 1994-242402
PATENT INFORMATION:
                                                    19961203
                                                                          <--
APPLICATION INFO.:
                                                    19940513
                                                              (8)
DOCUMENT TYPE:
                          Utility
FILE SEGMENT:
                          Granted
PRIMARY EXAMINER:
                          LeGuyader, John L.
ASSISTANT EXAMINER:
                          Larson, Thomas G.
LEGAL REPRESENTATIVE:
                          Logan, April C.
```

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

15 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 3698

CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB

The present invention discloses nucleic acid enzymes capable of cleaving nucleic acid molecules, including single-stranded DNA, in a site-specific manner under physiologic conditions, as well as compositions including same. The present invention also discloses methods of making and using the disclosed enzymes and compositions.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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ANSWER 15 OF 27
                 USPATFULL on STN
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ACCESSION NUMBER:

96:106381 USPATFULL

INVENTOR(S):

TITLE:

Antisense oligonucleotide inhibition of the RAS gene Monia, Brett P., Carlsbad, CA, United States

Freier, Susan M., San Diego, CA, United States Ecker, David J., Leucadia, CA, United States

PATENT ASSIGNEE(S): Isis Pharmaceuticals Inc., Carlsbad, CA, United States NUMBER KIND DATE

US 5576208 US 1994-297248 PATENT INFORMATION: 19961119 <--APPLICATION INFO.:

19940826 (8) RELATED APPLN. INFO.:

Continuation of Ser. No. US 1993-7996, filed on 21 Jan 1993, now abandoned which is a continuation-in-part of Ser. No. US 1991-715196, filed on 14 Jun 1991, now abandoned And a continuation-in-part of Ser. No. US 1992-958134, filed on 5 Oct 1992, now abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER:

Elliott, George C. Law Offices of Jane Massey Licata LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1,3

NUMBER OF DRAWINGS: 26 Drawing Figure(s); 21 Drawing Page(s)

LINE COUNT: 1666

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Compositions and methods are provided for the modulation of expression AB

of the human ras gene in both the normal and activated forms.

Oligonucleotides are provided which are specifically hybridizable with

RNA or DNA deriving from the human ras gene, having nucleotide
units sufficient in identity and number to effect such specific
hybridization. Oligonucleotides specifically hybridizable with a translation initiation site or with the codon-12 mutation of activated ras are provided. Such oligonucleotides can be used for diagnostics as well as for research purposes. Methods are also disclosed for modulating ras gene expression in cells and tissues using the oligonucleotides provided, and for specific modulation of expression of the activated ras gene. Methods for diagnosis, detection and treatment of conditions arising from the activation of the H-ras gene are also disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 16 OF 27 USPATFULL ON STN ACCESSION NUMBER:

96:97025 USPATFULL TITLE: Texaphyrins and uses thereof

INVENTOR(S):

Magda, Darren, Cupertino, CA, United States Sessler, Jonathan L., Austin, TX, United States Iverson, Brent, Austin, TX, United States

Jansen, Petra L., Austin, TX, United States Wright, Meredith, San Jose, CA, United States Mody, Tarak D., Sunnyvale, CA, United States Hemmi, Gregory W., Sunnyvale, CA, United States University of Texas, Austin, TX, United States (U.S.

PATENT ASSIGNEE(S): corporation)

Pharmacyclics, Inc., Sunnyvale, CA, United States (U.S.

corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5567687 APPLICATION INFO.:

RELATED APPLN. INFO.:

19961022 us 1994-310501 19940921 (8)

Continuation-in-part of Ser. No. US 1993-112872, filed on 25 Aug 1993, now patented, Pat. No. US 5451576 And Ser. No. US 1994-227370, filed on 14 Apr 1994 which is a continuation-in-part of Ser. No. US 1993-75123, filed

on 9 Jun 1993, now abandoned which is a

continuation-in-part of Ser. No. US 1992-822964, filed on 21 Jan 1992, now patented, Pat. No. US 5252720, issued on 12 Oct 1993 which is a continuation-in-part of Ser. No. US 1991-771393, filed on 30 Sep 1991, now abandoned which is a continuation-in-part of Ser. No. US 1990-539975, filed on 18 Jun 1990, now patented, Pat. No. US 5162509, issued on 10 Nov 1992 which is a division of Ser. No. US 1989-320293, filed on 6 Mar 1989, now patented, Pat. No. US 4935498, issued on 19

Jun 1990 , said Ser. No. US -112872 which is a

division of Ser. No. US -822964

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Raymond, Richard L. LEGAL REPRESENTATIVE: Arnold, White & Durkee

NUMBER OF CLAIMS: 13 **EXEMPLARY CLAIM:**

LINE COUNT: 2828

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A texaphyrin having substituents containing ethoxy groups, methods for using texaphyrins in photodynamic therapy, and cleavage of a polymer of deoxyribonucleic acid are disclosed. The in vivo treatment of tumors and atheroma is demonstrated using Lu(III)texaphyrin complexes. A preferred method of use is the site-specific cleavage of a polymer of deoxyribonucleic acid and a preferred texaphyrin is a derivatized texaphyrin having binding specificity, in particular, a texaphyrin covalently coupled to a site-directing molecule, preferably an oligonucleotide.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 17 OF 27 USPATFULL on STN

ACCESSION NUMBER: 96:43784 USPATFULL

TITLE: Pyrrolidine-containing monomers and oligomers Acevedo, Oscar L., San Diego, CA, United States
Hebert, Normand, San Marcos, CA, United States
Isis Pharmaceuticals, Inc., Carlsbad, CA, United States INVENTOR(S):

PATENT ASSIGNEE(S):

(U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: us 5519134 19960521 <--

US 1994-180134 APPLICATION INFO.: 19940111 (8)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: McKane, Joseph K.

Woodcock Washburn Kurtz Mackiewicz & Norris LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: 16 **EXEMPLARY CLAIM:**

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 1864

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Novel pyrrolidine monomers bearing various functional groups are used to AB prepare oligomeric structures. The pyrrolidine monomers can be joined via standard phosphate linkages including phosphodiester and phosphorothioate linkages. Useful functional groups include nucleobases

as well as polar groups, hydrophobic groups, ionic groups, aromatic

groups and/or groups that participate in hydrogen-bonding.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 18 OF 27 USPATFULL on STN

ACCESSION NUMBER: 96:41104 USPATFULL

Enzymatic synthesis of repeat regions of TITLE:

oligonucleotides

INVENTOR(S): Hyman, Edward D., 2100 Sawmill Rd. Apt. 4-103. River

Ridge, LA, United States 70123

NUMBER KIND DATE

US 5516664 US 1993-161224 PATENT INFORMATION: 19960514

APPLICATION INFO.: 19931202 (8)

Continuation-in-part of Ser. No. US 1993-100671, filed RELATED APPLN. INFO.: on 30 Jul 1993 which is a continuation-in-part of Ser. No. US 1992-995791, filed on 23 Dec 1992, now patented,

Pat. No. US 5436143

Utility DOCUMENT TYPE: FILE SEGMENT: Granted PRIMARY EXAMINER: Naff, David M. ASSISTANT EXAMINER: Prats, Francisco C. LEGAL REPRESENTATIVE: Oppedahl & Larson

NUMBER OF CLAIMS: 21 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 5 Drawing Figure(s); 5 Drawing Page(s)

LINE COUNT: 1099

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Enzymatic synthesis of a repeat region of an oligonucleotide may be performed by the steps of: (a) combining a primer and a blocked nucleotide in the presence of a chain extending enzyme whereby a primer-blocked nucleotide product is formed containing the blocked nucleotide coupled to the primer at its 3'-end; (b) removing the blocking group from the 3'-end of the primer-blocked nucleotide product using a 3'-phosphatase enzyme substantially without removing the

nucleotide; and (c) repeating the cycle of steps (a) and (b), using the primer-nucleotide product of step (b) as the primer for step (a) in the next cycle, for sufficient cycles to form the oligonucleotide product. These cycles are performed preferably in a single vessel without intermediate purification of oligonucleotide product.

Also disclosed is a process for synthesizing an oligonucleotide having a defined sequence including at least one repeat region and one non-repeating region, wherein at least one non-repeating region is synthesized by reaction cycles using the steps of extending a primer with a 3'-blocked nucleotide, inactivating unreacted 3'-blocked nucleotide, and removing the blocking group from the extended primer. The disclosed processes may be used to synthesize repeat regions of The disclosed processes may be used to synthesize repeat regions of oligoribonucleotides.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 19 OF 27 USPATFULL ON STN SSION NUMBER: 96:38773 USPATFULL

ACCESSION NUMBER:

TITLE:

Stem-loop_oligonucleotides containing parallel and

antiparallel binding domains

INVENTOR(S):

Kool, Eric T., Rochester, NY, United States

Research Corporation Technologies, Inc., Tucson, AZ, PATENT ASSIGNEE(S):

United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION:

US 5514546 19960507 <--

APPLICATION INFO.: DOCUMENT TYPE:

US 1993-115497 19930901 (8)

FILE SEGMENT:

Utility Granted

PRIMARY EXAMINER:

Jones, W. Gary

ASSISTANT EXAMINER: LEGAL REPRESENTATIVE: Myers, Carla Scully, Scott, Murphy & Presser

NUMBER OF CLAIMS: **EXEMPLARY CLAIM:**

NUMBER OF DRAWINGS:

8 Drawing Figure(s); 8 Drawing Page(s)

2331 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT. AΒ

The present invention provides stem-loop oligonucleotides containing a The present invention provides stem-loop origonucleotides containing a double-stranded stem domain of at least about 2 base pairs and a single-stranded loop domain. The loop domains of the present oligonucleotides include at least one parallel binding (P) domain separated by at least about 3 nucleotides from a corresponding anti-parallel binding (AP) domain. Each P and corresponding AP domain of the present oligonucleotides can bind detectably to one strand of a defined nucleic acid target wherein the P domain binds in a parallel manner to the target and the corresponding AP domain binds in an anti-parallel manner to the target. The present stem-loop oligonucleotides can bind to both single-stranded and double-stranded target nucleic acids. The present invention also provides methods of using these oligonucleotides as well as kits and pharmaceutical compositions containing these oligonucleotides.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 20 OF 27 USPATFULL on STN

ACCESSION NUMBER:

96:36439 USPATFULL

TITLE:

RNA Inhibiting expression by forming a pseudo-half-knot ***RNA*** at the target's

INVENTOR(S):

PATENT ASSIGNEE(S):

Ecker, David, Leucadia, CA, United States
ISIS Pharmaceuticals, Inc., Carlsbad, GA, United States

(U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: APPLICATION INFO.:

US 5512438 19960430 US 1994-176314 19940103 (8)

RELATED APPLN. INFO.:

Continuation of Ser. No. US 1992-916764, filed on 20

Jul_1992, now abandoned DOCUMENT TYPE:

Utility

FILE SEGMENT: PRIMARY EXAMINER:

Granted Jones, W. Gary Tran, Paul B.

ASSISTANT EXAMINER: LEGAL REPRESENTATIVE:

Woodcock Washburn Kurtz MacKiewicz & Norris

NUMBER OF CLAIMS:

NUMBER OF DRAWINGS: 18 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 908

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

EXING IS AVAILABLE FUR THIS PATENT.

Compositions and methods for modulating the activity of ***RNA**

Compositions and methods for modulating the activity of ***RNA*** Compositions and methods for moderating the standard with an ***RNA** are provided. Oligonucleotides are hybridized with an ***RNA*** longer recognized by its regulatory protein after oligonucleotide binding. Reactive moieties can be tethered to the oligonucleotide that enhance its activity. Antisense oligonucleotides directed against HIV TAR are provided.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 21 OF 27 USPATFULL on STN

96:19211 USPATFULL ACCESSION NUMBER:

Nucleic acid ligands to HIV-RT and HIV-1 rev TITLE: INVENTOR(S): Gold, Larry, Boulder, CO, United States

Tuerk, Craig, Morehead, KY, United States

PATENT ASSIGNEE(S): NeXstar Pharmaceuticals, Inc., Boulder, CO, United

States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: us 5496938 19960305

APPLICATION INFO.: us 1992-964624 19921021 (7)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1991-714131, filed

on 10 Jun 1991 And a continuation-in-part of Ser. No. 1990-536428, filed on 11 Jun 1990, now abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Zitomer, Stephanie W. LEGAL REPRESENTATIVE: Swanson & Bratschun

NUMBER OF CLAIMS: 10 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 44 Drawing Figure(s); 34 Drawing Page(s)

LINE COUNT: 2438

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Methods for the identification and production of improved nucleic acid AB ligands are based on the SELEX process. Nucleic acid ligands to HIV-RT and HIV-1 Rev are identified according to the methods described herein.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 22 OF 27 MEDLINE on STN DUPLICATE 1

96196016 ACCESSION NUMBER: MEDLINE PubMed ID: 8608452 DOCUMENT NUMBER:

RNase H cleavage for processing of in vitro transcribed TITLE: ***RNA*** for NMR studies and ***RNA*** ligation.

Lapham J; Crothers D M **AUTHOR:**

CORPORATE SOURCE: Department of Chemistry, Yale University, New Haven,

Connecticut 06511, USA.

GM-21966 (NIGMS) CONTRACT NUMBER:

RNA (New York, N.Y.), ***(1996 Mar)**
Journal code: 9509184. ISSN: 1355-8382. SOURCE: ***(1996 Mar)*** 2 (3) 289-96.

PUB. COUNTRY: United States

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199605

ENTRY DATE: Entered STN: 19960605

Last Updated on STN: 19990129

Entered Medline: 19960528 es of ***RNA*** for st AB Large quantities of for study by NMR and X-ray crystallography can be produced by transcription reactions in vitro using T7 bacteriophage ***RNA*** polymerase. A limitation on producing ***RNA*** with this polymerase has been the strong dependence of the yield of the transcription reaction on the sequence at the 5' end of the produced. We report a procedure for obtaining large enzymatically synthesized ***RNA*** from T7 ***RNA*** quantities of enzymatically synthesized polymerase that has no dependence on the 5' end sequence of the target ***RNA*** . Ribonuclease H has been shown previously (Inoue H, Hayase Y, Iwai S, Ohtsuka E, 1987, FEBS Lett 215:327-330) to cleave ***RNA*** site specifically using ***2*** '- ***0*** - ***methyl*** site specifically using ***RNA*** /DNA Chimeras to direct the cleavage site. We show that
2 '- ***O*** - ***methyl*** ***RNA*** nucleotides or

5'-side of the DNA nucleotides in the chimera are not essential for

RNA nucleotides on the

the same ***2*** '- ***0*** - ***methyl*** chimera may be used to process any ***RNA*** sequence. We have adapted this reaction to the cleavage of NMR-scale quantities of ***RNA*** at high yield.

RNA is synthesized using T7 ***RNA*** polymerase with a 15-nt high-yielding leader sequence at the 5' end, and then this sequence is cleaved off with the RNase H cleavage reaction. The cleaved ***RNA*** has 3'-hydroxyl and 5'-phosphate ends, so that the products can be used directly as substrates for ligation by T4 DNA ligase. We show that the cleavage reaction occurs efficiently in solution and on a solid streptavidin/agarose matrix. We report an example in which we are able to improve transcription yield by more than five-fold using this technique in the synthesis of a 15N isotopically labeled ***hairpin*** found in the Crithidia fasciculata spliced leader ***RNA***. We are able to obtain a 0.5-mM NMR sample from this inherently poorly transcribing sequence, while minimizing the amount of isotopically labeled rNTPs used to produce it. The NMR spectroscopic results are consistent with the predicted ***RNA***

it. The NMR spectroscopic results are consistent with the predicted ***RNA*** secondary structure. ANSWER 23 OF 27 MEDLINE on STN DUPLICATE 2 ACCESSION NUMBER: 1998044723 MEDLINE DOCUMENT NUMBER: PubMed ID: 9383475 TITLE: Nuclease-resistant nucleic acid ligands to vascular permeability factor/vascular endothelial growth factor. Green L S; Jellinek D; Bell C; Beebe L A; Feistner B D; **AUTHOR:** Gill S C; Jucker F M; Janjic N NeXstar Pharmaceuticals, Boulder, Colorado 80301, USA. Chemistry & biology, ***(1995_Oct)*** 2 (10) 683-9 CORPORATE SOURCE: SOURCE: 2 (10) 683-95. Journal code: 9500160. ISSN: 1074-5521. PUB. COUNTRY: ENGLAND: United Kingdom Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE: English LANGUAGE: FILE SEGMENT: Priority Journals ENTRY MONTH: 199801 Entered STN: 19980129 **ENTRY DATE:** Last Updated on STN: 19980129 Entered Medline: 19980115

BACKGROUND: Vascular permeability factor/vascular endothelial growth factor (VPF/VEGF) is a potent inducer of new blood vessel growth (angiogenesis) that contributes to the pathology of many angiogenesis-associated disease states such as psoriasis, rheumatoid arthritis and cancer. Few molecular entities capable of binding to AΒ VPF/VEGF with high affinity and specificity have been described to date.

RESULTS: Nuclease-resistant 2'-amino-2'-deoxypyrimidine nucleotide

RNA (2'-aminopyrimidine ***RNA***) ligands that bind to VPF/VEGF with high affinity have been identified by iterative rounds of affinity-selection/amplification from two independent random libraries. The sequence information that confers high affinity binding to VPF/VEGF is contained in a contiguous stretch of 24 nucleotides, 5'CCCUGAUGGUAGACGCCGGGGUG-3' (2'-aminopyrimidine nucleotides are designated with italic letters). Of the 14 ribopurines in this minimal ligand, 10 can be substituted with the corresponding 2'-0-methylpurine nucleotides without a reduction in hinding affinity to VPE/VEGE. In fact, the without a reduction in binding affinity to VPF/VEGF. In fact, the ***2*** '- ***O*** - ***methyl*** substitution at permissi substitution at permissive positions leads to a approximately 17-fold improvement in the binding affinity to VPF/VEGF. The higher affinity results from the reduction in the dissociation rate constant of the ***2*** '- ***0*** - ***methyl*** -substituted ***RNA*** ligand from the protein compared to the unsubstituted ligand. The ***2*** '- ***0*** - ***methyl*** -substituted minimal ligand, which folds into a bulged ***hairpin*** motif is also more thermally stable than the unsubstituted ligand motif, is also more thermally stable than the unsubstituted ligand. Nuclease resistance of the ligand is further improved by the ***O*** - ***methyl*** substitutions and the addition of ***O*** - ***methyl*** substitutions and the addition of short phosphorothioate caps to the 3'- and 5'-ends. CONCLUSIONS: We have used the SELEX (systematic evolution of ligands by exponential enrichment) process in conjunction with post-SELEX modifications to define a highly nuclease-resistant oligonucleotide that binds to VPF/VEGF with high affinity and specificity.

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L4 ANSWER 24 OF 27 MEDLINE ON STN
ACCESSION NUMBER: 96113282 MEDLINE
DOCUMENT NUMBER: Pubmed ID: 8785472
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TITLE: 4'-Thio- ***RNA*** : synthesis of mixed base

4'-thio-oligoribonucleotides, nuclease resistance, and base pairing properties with complementary single and double

strand.

AUTHOR: Leydier C; Bellon L; Barascut J L; Morvan F; Rayner B;

Laboratoire de Chimie Bio-Organique, URA 488, CNRS, Universite de Montpellier II, France. CORPORATE SOURCE:

(1995 Fall) Antisense research and development, SOURCE:

(3) 167-74.

Journal code: 9110698. ISSN: 1050-5261.

PUB. COUNTRY: **United States**

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199609

ENTRY DATE: Entered STN: 19961008

Last Updated on STN: 19961008 Entered Medline: 19960926

AB 4'-Thio-beta-D-oligoribonucleotides (12 mer and 16 mer) containing a mixed base sequence were synthesized via the phosphoramidite solid support approach. These ***RNA*** analogs showed very good nuclease _analogs showed very good nuclease resistance as compared with wild-type ***RNA*** . Furthermore, 4'-thio-beta-D-oligoribonucleotides were shown to hybridize with a complementary DNA or ***RNA*** strand to form a duplex or with a DNA ***hairpin*** to form a triple helix. 4'-Thio- ***RNA*** binds most tightly to its complementary ***RNA*** strand than to its complementary DNA strand. A 4'-thio- ***RNA*** : ***RNA*** duplex as stable as a ***2*** '- ***0*** - ***methyl*** - ***RNA*** : binds more duplex is duplex. 4'-Thio- ***RNA*** , however, forms a 4'-thio-***RNA*** :DNA:DNA triplex with a stability similar to the corresponding

ANSWER 25 OF 27 MEDLINE on STN **DUPLICATE 4**

ACCESSION NUMBER: 93374933 MEDLINE DOCUMENT NUMBER: PubMed ID: 7690032

triplex with all wild-type DNA.

Four ribose 2'-hydroxyl groups essential for catalytic function of the ***hairpin*** ribozyme. TITLE:

Chowrira B M; Berzal-Herranz A; Keller C F; Burke J M Markey Center for Molecular Genetics, Department of AUTHOR: CORPORATE SOURCE:

Microbiology and Molecular Genetics, University of Vermont,

Burlington 05405.

Journal of biological chemistry,
268 (26) 19458-62. SOURCE: ***(1993 Sep 15)***

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199310

ENTRY DATE:

Entered STN: 19931022 Last Updated on STN: 19980206 Entered Medline: 19931007

hairpin AB ribozyme catalyzes site-specific cleavage of an substrate using a magnesium-dependent transphosphorylation mechanism. Here, we describe experiments designed to test the importance of ribose 2'-hydroxyl groups for ribozyme function. Ribozymes for this work were synthesized in two segments using solid-phase ***RNA*** work were synthesized in two segments using solid-phase ***RNA***
phosphoramidite chemistry. 2'-Deoxyribonucleotides were systematically introduced at each of the 50 positions within the ribozyme, and the catalytic activity of the resulting mixed ***RNA*** -DNA polymers was measured. Deletion of the 2'-hydroxyl group at each of four sites (A10, G11, A24, and C25) was found to result in severe inhibition of cleavage activity (kcat/kM decreased by 100- to 1000-fold), although KM ***RNA*** -DNA polymers was measurements and mobility-shift assays showed that substrate binding was not affected. Identical results were obtained upon substitution of these ribonucleotides with ***2*** '- ***0*** - ***methyl*** derivatives. Inhibition by 2'-modified sugars at G11 or A24 was rescued by increased Mg2+ concentrations, suggesting that these 2'-hydroxyls may function in magnesium binding. Our results demonstrate that the 2'-hydroxyl groups at A10, G11, A24, and C25 provide essential functions for catalysis, possibly forming important tertiary contacts or magnesium coordination sites that are necessary for active site architecture.

ANSWER 26 OF 27 MEDLINE on STN DUPLICATE 5

ACCESSION NUMBER: 92375676 MEDLINE DOCUMENT NUMBER: PubMed ID: 1508680

TITLE: Enhancement of ribosomal frameshifting by oligonucleotides

targeted to the HIV gag-pol region. Vickers T A; Ecker D J

AUTHOR:

CORPORATE SOURCE: ISIS Pharmaceuticals, Carlsbad, CA 92008 Nucleic acids research, ***(1992 Aug 11)*** SOURCE: 20 (15) Journal code: 0411011. ISSN: 0305-1048.

ENGLAND: United Kingdom PUB. COUNTRY:

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals; AIDS ENTRY MONTH: 199209

ENTRY DATE: Entered STN: 19921009

Last Updated on STN: 19970203

Entered Medline: 19920923

The pol gene of all retroviruses is expressed as a gag-pol fusion protein which is proteolytically processed to produce all viral enzymes. In the human immunodeficiency virus (HIV), the gag and pol genes overlap by 241 nucleotides with pol in the -1 phase with respect to gag. The gag-pol fusion is produced via a -1 ribosomal frameshifting event that brings the AB overlapping, out-of-phase gag and pol genes into translational phase.

Frameshifting occurs at a so called 'shift site' 8-10 nucleotides upstream of a ***hairpin*** loop which may play a role in the regulation of frameshifting. We have fused this region of HIV-1 to the 5' end of the firefly luciferase reporter gene in order to quantitatively measure ribosomal frameshifting both in cells and by in vitro translation. A series of ***2*** '- ***0*** - ***methyl*** oligonucleotides was designed to specifically hind the sequences which flank the gag-pol designed to specifically bind the sequences which flank the gag-pol ***hairpin*** . Ribosomal frameshifting is enhanced up to 6 fold by those oligonucleotides which bind the area just 3 to the stem.
Oligonucleotides which bind 5' to the stem have no effect on frameshift efficiency. In addition, we have constructed a series of fusion genes which mimic the effect of the bound oligonucleotides with intramolecular hairpins. The results suggest that increasing ***RNA*** secondary structure downstream of the shift site increases the frequency of ribosomal frameshifting, and that this effect can be mimicked by antisense oligonucleotides oligonucleotides.

L4 ANSWER 27 OF 27 MEDLINE on STN **DUPLICATE 6**

84169564 ACCESSION NUMBER: MEDLINE DOCUMENT NUMBER: PubMed ID: 6424099

RNA : experimental TITLE: Xenopus laevis 18S ribosomal

determination of secondary structural elements, and locations of methyl groups in the secondary structure

model

Atmadja J; Brimacombe R; Maden B E AUTHOR:

SOURCE: Nucleic acids research. ***(1984 Mar 26)*** 12 (6)

2649-67.

Journal code: 0411011. ISSN: 0305-1048.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198405

ENTRY DATE: Entered STN: 19900319

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RNA 18S ribosomal AB from X. laevis was subjected to partial digestion with ribonucleases A or T1 under a variety of conditions, and base-paired fragments were isolated. Sequence analysis of the fragments enabled five base-paired secondary structural elements of the 18S ***RNA*** to be established. Four of these elements (covering bases 221-256, 713-757, 1494-1555 and 1669-1779) confirm our previous secondary structure predictions, whereas the fifth (comprising bases 1103-1125) represents a phylogenetically conserved "switch" structure, which can also form in prokaryotic 16S ***RNA*** The results are incorporated into a refined model of the 18S ***RNA*** secondary structure, which also includes the locations of the many methyl groups in Y large 185 includes the locations of the many methyl groups in X. laevis 18S ***RNA*** . In general the methyl groups occur in non-helical regions, at ***hairpin*** loop ends, or at helix boundaries and imperfections. One large cluster of ***2*** '- ***0*** - ***methyl*** groups One large cluster of ***2*** '- ***O*** - ***methyl*** groups occurs in a region of complicated secondary structure in the 5'-one third